



Serial No. 09/457,066

RECEIVED  
JUN 27 2001  
TECH CENTER 1600/2900

### Appendix

3. (amended) An isolated polypeptide which is from 111 to 136 amino acid residues in length and comprises a sequence of amino acid residues as shown in SEQ ID NO:2 from residue 235 to residue 345 [comprising a sequence of amino acids of the formula  $R1_x-R2_y-R3_z$ , wherein:

R1 comprises a polypeptide of from 100 to 120 residues in length that is at least 90% identical to residues 46-163 of SEQ ID NO:2, and comprises a sequence motif C[KR]Y[DNE][WYF]X(11,15)G[KR][WYF]C (SEQ ID NO:4) corresponding to residues 104-124 of SEQ ID NO:2;

R2 is a polypeptide at least 90% identical to residues 164-234 of SEQ ID NO:2;

R3 is a polypeptide at least 90% identical in amino acid sequence to residues 235-345 of SEQ ID NO:2 and comprises cysteine residues at positions corresponding to residues 250, 280, 284, 296, 335, and 337 of SEQ ID NO:2; a glycine residue at a position corresponding to residue 282 of SEQ ID NO:2; and a sequence motif CX(18,33)CXGXCX(6,33)CX(20,40)CXC (SEQ ID NO:3) corresponding to residues 250-337 of SEQ ID NO:2; and

each of x, y, and z is individually 0 or 1, subject to the limitations that:

at least one of x and z is 1; and

if x and z are each 1, then y is 1].

11. (amended) The isolated polypeptide of claim [10] 3 comprising a sequence of amino acid residues as shown in [wherein R3 comprises residues 235-345 of] SEQ ID NO:2 from residue 230 to residue 345.

17. (amended) An isolated protein comprising a first polypeptide [operably linked] disulfide bonded to a second polypeptide, wherein each of said first and second polypeptides is from 111 to 136 amino acid residues in length and comprises a sequence of amino acid residues as shown in SEQ ID NO:2 from residue 235 to residue 345 [polypeptide comprises a sequence of amino acids of the formula  $R1_x-R2_y-R3_z$ , wherein:

R1 comprises a polypeptide of from 100 to 120 residues in length that is at least 90% identical to residues 46-163 of SEQ ID NO:2, and comprises a sequence

A

motif C[KR]Y[DNE][WYF]X{11,15}G[KR][WYF]C (SEQ ID NO:4) corresponding to residues 104-124 of SEQ ID NO:2;

R2 is a polypeptide at least 90% identical to residues 164-234 of SEQ ID NO:2;

R3 is a polypeptide at least 90% identical in amino acid sequence to residues 235-345 of SEQ ID NO:2 and comprises cysteine residues at positions corresponding to residues 250, 280, 284, 296, 335, and 337 of SEQ ID NO:2; a glycine residue at a position corresponding to residue 282 of SEQ ID NO:2; and a sequence motif CX{25,33}CXGXCX{10,33}CX{20,40}CXC (SEQ ID NO:3) corresponding to residues 250-337 of SEQ ID NO:2; and

each of x, y, and z is individually 0 or 1, subject to the limitations that:

at least one of x and z is 1; and

if x and z are each 1, then y is 1,

and wherein said protein modulates cell proliferation, differentiation, metabolism, or migration.

22. (amended) The isolated protein according to claim [21] 17 wherein each of said first and second polypeptides comprises [residues 235-345 of] a sequence of amino acid residues as shown in SEQ ID NO:2 from residue 230 to residue 345.

25. (amended) An isolated polynucleotide [of up to approximately 4 kb in length, wherein said polynucleotide encodes] encoding a polypeptide which is from 111 to 136 amino acid residues in length and comprises a sequence of amino acid residues as shown in SEQ ID NO:2 from residue 235 to residue 345 [comprising a sequence of amino acids of the formula R1<sub>x</sub>-R2<sub>y</sub>-R3<sub>z</sub>, wherein:

R1 comprises a polypeptide of from 100 to 120 residues in length that is at least 90% identical to residues 46-163 of SEQ ID NO:2, and comprises a sequence motif C[KR]Y[DNE][WYF]X{11,15}G[KR][WYF]C (SEQ ID NO:4) corresponding to residues 104-124 of SEQ ID NO:2;

R2 is a polypeptide at least 90% identical to residues 164-234 of SEQ ID NO:2;

R3 is a polypeptide at least 90% identical in amino acid sequence to residues 235-345 of SEQ ID NO:2 and comprises cysteine residues at positions corresponding to residues 250, 280, 284, 296, 335, and 337 of SEQ ID NO:2; a glycine residue at a position corresponding to residue 282 of SEQ ID NO:2; and a sequence

A

motif CX{25,33}CXGXCX{10,33}CX{20,40}CXC (SEQ ID NO:3) corresponding to residues 250-337 of SEQ ID NO:2; and

each of x, y, and z is individually 0 or 1, subject to the limitations that:

at least one of x and z is 1; and

if x and z are each 1, then y is 1].

32. (amended) A method of producing a protein comprising:

culturing a cell into which has been introduced an expression vector according to claim 28, whereby said cell expresses the polypeptide encoded by the DNA segment; and

recovering a protein comprising the expressed [protein] polypeptide.

A